

That which is claimed is:

1. A composition comprising the ligand binding domain of a farnesoid X receptor (FXR) in crystalline form.
2. A composition according to claim 1 further comprising a ligand of said FXR.
3. A composition according to claim 2, wherein said ligand is selected from the group consisting of fexaramine, fexarine, fexarene and GW4064.
4. A composition according to claim 2, wherein said ligand is selected from the group consisting of fexaramine, fexarine and fexarene.
5. A composition according to claim 2, wherein said ligand is fexaramine.
6. A composition according to claim 1 as described by the structure coordinates set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.
7. A composition according to claim 2 as described by the structure coordinates set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and said ligand.
8. A composition according to claim 2, wherein the crystals belong to space group  $P2_12_12_1$  with unit cell dimensions of about:  
 $a = 37 \text{ \AA}$ ,  $b = 57 \text{ \AA}$ ,  $c = 117 \text{ \AA}$ ,  
 $\alpha = 90^\circ$ ,  $\beta = 90^\circ$ , and  $\gamma = 90^\circ$ .
9. A composition according to claim 2, wherein the crystals belong to space group  $P2_12_12_1$  with unit cell dimensions:  
 $a = 36.656 \text{ \AA}$ ,  $b = 56.776 \text{ \AA}$ ,  $c = 117.646 \text{ \AA}$ ,  
 $\alpha = 90.0^\circ$ ,  $\beta = 90.0^\circ$ , and  $\gamma = 90.0^\circ$ .

10. A composition according to claim 1, wherein said ligand binding domain comprises amino acid residues 248 – 476 of SEQ ID NO:1.

11. A computer for producing a three-dimensional representation of a farnesoid X receptor (FXR) molecule or molecular complex or a homologue of said FXR molecule or molecular complex, wherein said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex comprises a ligand binding domain defined by structure coordinates obtained from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex, said computer comprising:

- (i) a computer-readable data storage medium comprising a data storage material encoded with computer-readable data, wherein said data comprises X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex;
- (ii) a working memory for storing instructions for processing said computer-readable data;
- (iii) a central-processing unit coupled to said working memory and to said computer-readable data storage medium for processing said computer-machine readable data into said three-dimensional representation; and
- (iv) a display coupled to said central-processing unit for displaying said three-dimensional representation.

12. A computer according to claim 11, wherein said structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.

13. A computer for determining at least a portion of the structure coordinates corresponding to X-ray diffraction data obtained from a farnesoid X receptor (FXR) molecule or molecular complex or a homologue of said FXR molecule or molecular complex, said computer comprising:

(i) a computer-readable data storage medium comprising a data storage material encoded with computer-readable data, wherein said data comprises at least a portion of the structure coordinates of Appendix 1;

(ii) a computer-readable data storage medium comprising a data storage material encoded with computer-readable data, wherein said data comprises X-ray diffraction data obtained from said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex;

(iii) a working memory for storing instructions for processing said computer-readable data of (i) and (ii);

(iv) a central-processing unit coupled to said working memory and to said computer-readable data storage medium of (i) and (ii) for performing a Fourier transform of the machine readable data of (i) and for processing said computer-readable data of (ii) into structure coordinates; and

(v) a display coupled to said central-processing unit for displaying said structure coordinates of said FXR molecule or molecular complex.

14. A method of predicting a molecule capable of binding to a farnesoid X receptor (FXR) molecule, said method comprising:

modeling a test molecule that potentially interacts with the ligand binding domain of said FXR molecule, wherein said ligand binding domain is defined by a plurality of structure coordinates of the ligand binding domain of a FXR molecule or a fragment thereof,

wherein said structure coordinates are derived from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex.

15. A method according to claim 14, wherein said plurality of structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.
16. A method according to claim 14, wherein said test molecule is developed de novo.
17. A method according to claim 14, wherein said test molecule is developed from a known ligand of FXR.
18. A method according to claim 14, wherein said test molecule is developed using a computer algorithm to predict a three-dimensional representation of said test molecule interacting with a FXR based upon a three-dimensional representation of a FXR molecule or fragment thereof.
19. A method of identifying a compound with agonist activity for a farnesoid X receptor (FXR) molecule, said method comprising:
  - (a) modeling a test compound that potentially interacts with the ligand binding domain of said FXR molecule or a fragment thereof, wherein said ligand binding domain is defined by a plurality of structure coordinates of the ligand binding domain of a FXR molecule or a fragment thereof,  
wherein said plurality of structure coordinates are derived from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex; and
  - (b) determining the ability of said test compound to activate said FXR molecule.
20. A method according to claim 19, wherein said plurality of structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.

21. A compound identified by the method of claim 19.
22. A pharmaceutical composition comprising a compound identified by the method of claim 19 and a pharmaceutically acceptable carrier therefor.
23. A method of identifying a compound with antagonist activity for a farnesoid X receptor (FXR) molecule, said method comprising:
- (a) modeling a test compound that potentially interacts with the ligand binding domain of said FXR molecule or a fragment thereof, wherein said ligand binding domain is defined by a plurality of structure coordinates of the ligand binding domain of a FXR molecule or a fragment thereof,  
wherein said plurality of structure coordinates are derived from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex; and
  - (b) determining the ability of said test compound to modulate the activity of said FXR molecule in the presence of a known FXR agonist.
24. A method according to claim 23, wherein said plurality of structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.
25. A compound identified by the method of claim 23.
26. A pharmaceutical composition comprising a compound identified by the method of claim 23 and a pharmaceutically acceptable carrier therefor.
27. A method of identifying a compound with partial agonist activity for a farnesoid X receptor (FXR) molecule, said method comprising:
- (a) modeling a test compound that potentially interacts with the ligand binding domain of said FXR molecule or a fragment thereof, wherein said ligand

binding domain is defined by a plurality of structure coordinates of the ligand binding domain of a FXR molecule or a fragment thereof,

wherein said plurality of structure coordinates are derived from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex; and

(b) determining the ability of said test compound to modulate the activity of said FXR molecule in the optional presence of a known FXR agonist.

28. A method according to claim 27, wherein said plurality of structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and a ligand therefor.

29. A compound identified by the method of claim 27.

30. A pharmaceutical composition comprising a compound identified by the method of claim 27 and a pharmaceutically acceptable carrier therefor.

31. A method for determining whether a test compound is capable of binding to the ligand binding domain of a farnesoid X receptor (FXR) molecule, said method comprising:

(a) determining the points of interaction between the ligand binding domain of a FXR, and one or more known ligand(s) therefor; and

(b) analyzing said test compound to determine whether similar points of interaction exist between said test compound and said ligand binding domain.

32. A method according to claim 31, wherein step (a) utilizes a plurality of structure coordinates derived from X-ray diffraction data obtained from crystals of said FXR molecule or molecular complex or a homologue of said FXR molecule or molecular complex to define said points of interaction.

33. A method according to claim 32, wherein said structure coordinates are set forth in Appendix 1, or a portion thereof sufficient to define the points of interaction between said ligand binding domain and said ligand(s).

34. A method according to claim 31, wherein said test compound is a bile acid.

35. A method according to claim 31, wherein said bile acid is a natural bile acid.

36. A method according to claim 31, wherein said bile acid is a modified bile acid.

37. A method according to claim 31, wherein said bile acid is a synthetic bile acid.